```
00001
1 IN THE UNITED STATES DISTRICT COURT FOR THE
 2
              EASTERN DISTRICT OF ARKANSAS
 3
                    WESTERN DIVISION
5 HENRY W. BOERNER, Individually ) and as Administrator of the 6 Estate of MARY JANE BOERNER, ) Deceased,
 7
                                                        Plaintiffs,
                                   )
8
                                   )
                                                                          No. LR-C-
                                               VS.
                                                                        )
98-427
9
                                   )
                                       BROWN & WILLIAMSON TOBACCO
                                                                       )
10 COMPANY,
                                   )
                                                                        )
                   Defendant.
11
                                  )
12
13
14
                       DEPOSITION OF
                SANFORD H. BARSKY, M.D.
15
                LOS ANGELES, CALIFORNIA
16
17
               FRIDAY, JANUARY 14, 2000
18
19
20
21 ATKINSON-BAKER, INC. COURT REPORTERS
22 330 North Brand Boulevard, Suite 250 Glendale, California 91203
23 (818) 551-7300
24 REPORTED BY: LISA MICHAELS, RPR, CSR NO. 6361
25 FILE NO.: 9A004B8
00002
      IN THE UNITED STATES DISTRICT COURT FOR THE
1
 2
              EASTERN DISTRICT OF ARKANSAS
3
                     WESTERN DIVISION
4
                          - - -
5 HENRY W. BOERNER, Individually ) and as Administrator of the 6 Estate of MARY JANE BOERNER, ) Deceased,
7
                                   )
                                                        Plaintiffs,
                                                                       )
                                                                           No. LR-C-
8
                                   )
                                              vs.
                                                                        )
98-427
                                     BROWN & WILLIAMSON TOBACCO
9
                                   )
                                                                        )
10 COMPANY,
                                   )
                                                                        )
11
                   Defendant.
                                  )
12
13
14
15
16
          Deposition of SANFORD H. BARSKY, M.D., taken
17 on behalf of the Plaintiffs at 10422 Lindbrook
18 Drive, Los Angeles, California, commencing at 10:05
19 A.M., on Friday, January 14, 2000, before
20 Lisa Michaels, RPR, CSR No. 6361.
21
22
23
24
25
00003
                  APPEARANCES
1
 2
 3 FOR THE PLAINTIFFS:
 4
        SPOHRER, WILNER, MAXWELL & MATTHEWS BY: STEPHANIE HARTLEY,
ATTORNEY AT LAW
 5 444 East Duval Street Jacksonville, Florida 32202
 6
          (904) 354-8310
                                            - and-
 7
         GARY EUBANKS & ASSOCIATES
                                             BY: GERRY SCHULZE, ATTORNEY AT LAW
```

```
8
          708 West Second Street Little Rock, Arkansas 72201
         (501) 372-0266
9
10
11 FOR THE DEFENDANT:
    CHADBOURNE & PARKE BY: BRUCE G. SHEFFLER, ATTORNEY AT LAW
30 Rockefeller Plaza New York, New York 10112
12
13
          (212) 408-5100
                                         - and-
14
       DINSMORE & SHOHL BY: FRANK C. WOODSIDE, III, M.D.,
15
     ATTORNEY AT LAW 1900 Chemed Center 255 East Fifth Street Cincinnati, Ohio 45202 (513) 977-8266
16
17
18
19
20
21
22
23
2.4
2.5
00004
1
 2 WITNESS: SANFORD H. BARSKY, M.D.
 3 EXAMINATION
                                                PAGE
 4
         BY MS. HARTLEY
                                                 5
 5
         BY MR. SCHULZE
                                                 49
         BY MR. SHEFFLER
                                                 53
 6
7
8
9 EXHIBITS:
                      PLAINTIFFS' NUMBER
                                                        DESCRIPTION
10
PAGE
11 1 - Curriculum Vitae
12 2 - Handwritten notes "History"
13 and "Pathology Findings"
                                                     11
                                                    13
and "Pathology Findi
14 3 - "Code for Pictures"
                                                32
15 3-A through Photomicrographs
                                                       32
16 3-J
17 4 -
               Article entitled "Rising 53
                                                                    Incidence of
Bronchioloalveolar
    Lung Carcinoma and Its Unique Clinicopathologic
18
Features
19
20
21 QUESTIONS WITNESS WAS INSTRUCTED NOT TO ANSWER:
22
                         (NONE)
2.3
24 INFORMATION TO BE SUPPLIED:
25
                         (NONE)
00005
1 FRIDAY, JANUARY 14, 2000; LOS ANGELES, CALIFORNIA
                        10:05 A.M.
 3
 4
                 SANFORD H. BARSKY, M.D.,
 5
           having first been duly sworn, was
 6
            examined and testified as follows:
7
8
                      EXAMINATION
9 BY MS. HARTLEY:
10
    Q. Good morning, Dr. Barksy?
         A. Good morning.
Q. This is Stephanie Hartley. I believe
11
12
13 we have met before.
14 A. Well, we've never met except over
15 depositions.
```

```
Q.
               I think we met at a trial once.
16
17 Perhaps you don't recall. I was not the one asking
18 you the questions. But in any case, could you
19 please state your full name for the record,
20 please.
21
               Yes, it's Sanford H. Barksy, M.D.
          Α.
22
               And what is your professional address?
          Q.
              It's Department of Pathology, UCLA
23
          Α.
24 School of Medicine, Los Angeles, California,
25 90024.
00006
               And we're here today on the case of
1
2 Mary Jane Boerner; is that you're understanding?
 3
         A. Yes.
 4
          Q.
                When were you hired in this case,
5 Dr. Barksy?
6
         MR. SHEFFLER: Object to the form.
7
          THE WITNESS: I recall that in the summer
8 Bruce Sheffler came to see me with a bunch of
9 slides and asked my to review them, and after that
10 I received a set of medical records and that was in
11 the fall. So I guess you can say my "hiring" began
12 in the summer.
13 BY MS. HARTLEY:
14
       Q.
               And that was Bruce Sheffler who's with
15 you today?
16
      A. Yes.
              Do you know what company he
17
          Q.
18 represents?
    Α.
               You mean his law firm? Or his
19
20 client?
Q. No, his client.
22
               I think it's one of the tobacco
         Α.
23 companies. Brown & Williamson or something like
24 that.
25
              Have you testified for tobacco
          Q.
00007
1 companies in the past?
2 A. Yes.
3
          Ο.
               Can you tell me all of the cases in
4 which you've testified?
5
         A. There was one court case. There was
 6 one case that went to trial. It was a case of the
 7
   client's name was Conner.
8
          Q. Who was the tobacco company in that
9 case?
         A. I think that was R.J. Reynolds.Q. Have you testified in depositions in
10
11
12 any other tobacco cases?
13 A. Yes, there were -- I believe there
14 were two other cases that I've given depositions
15 other than the present one.
         Q. Correct.
16
17
          Α.
               There were two other cases.
         Q. Do you recall their names?A. You know, not offhand. If I thought
18
19
20 about it, they probably would come to me, but I
21 don't remember them right at this second.
        Q. Was one of them the Clark case?
22
23
          A. Yes, I believe so.
          Q.
               And who was the cigarette company in
25 that case?
80000
```

I'm not sure. I don't know at this Α. 2 point. I don't remember. Q. And in the third case you don't 3 4 remember the name of that case? 5 A. If you said the name to me, I might 6 recall it. Do you recall the cigarette company 7 Q. 8 involved in that case? A. I think the third case was R.J. 9 10 Reynolds. 11 Q. Have you ever been hired by lawyers 12 representing Brown & Williamson prior to this case? MR. SHEFFLER: Object to the form. 13 THE WITNESS: No. Well, I've looked at 14 15 slides on different cases, but it hasn't resulted 16 in a deposition. So again, it depends what you 17 mean by hired. 18 BY MS. HARTLEY: 19 Q. So you've looked at slides for Brown & 20 Williamson before this case? MR. SHEFFLER: Object to the form. 21 THE WITNESS: Bruce Sheffler over the past 22 23 ten years occasionally has brought me slides to 24 look at. So I assume that probably is that firm, 25 but I don't know for sure. 00009 1 BY MS. HARTLEY: Approximately how many cases has Bruce 3 Sheffler brought you over the past ten years? 4 MR. SHEFFLER: Again, let me just make an 5 objection for the form. THE WITNESS: Bruce Sheffler hasn't brought 6 7 me any cases in the last several years except for 8 this case, but let's say if we go back more than 9 five years -- I would say oh, maybe I've looked 10 at -- looked at 10 to 15 cases over the years, but 11 none recently. 12 BY MS. HARTLEY: 13 Approximately how many cases or sets Q. 14 of slides have you looked at for the tobacco 15 companies in the past ten years? MR. SHEFFLER: Again, let me object to the 16 17 form of the question. THE WITNESS: Oh, I would say 20 to 30 18 19 perhaps more. You know, many of these cases are 20 just a cursory slide review and that's it. So it's 21 hard for me to recount the exact number. 22 BY MS. HARTLEY: 23 Q. And when Bruce Sheffler brought you 24 these slides in this case, the Boerner case, back 25 in the summer of 1999, what did he ask you to do? 00010 Well, as he always asks me to do, he 1 Α. 2 starts off by asking me to review them in terms of, 3 you know, what pathology is present. So I review 4 them, look at them under the microscope, and that's 5 how it starts. 6 Q. And then what do you do? 7 Α. Well, it depends. Like I said, the 8 vast majority of these cases, that's last I hear of 9 them. In this particular case, he asked for my 10 opinion as to the pathology and asked me if I would 11 be willing to review the medical records, which I

12 said I would, and then he asked me, you know, what 13 my assessment of the case was. Q. Did he ask you about causation? 14 15 MR. SHEFFLER: Objection. Overbroad. THE WITNESS: We eventually got into that 16 17 area. So, yes, he did ask me about that. 18 BY MS. HARTLEY: 19 Q. Dr. Barsky, I have a C.V. that is not 20 up to date, but I just wanted to ask you if in the 21 past four years you have written any articles or 22 published any articles? Yes, I have. My C.V. changes, I would Α. 24 say, on a monthly basis, and I constantly update 25 it. I've given -- I've e-mailed Bruce Sheffler and 00011 1 given him a hard copy of an updated C.V. MR. SHEFFLER: Stephanie, if it makes it 2 3 easier, we can mark it as your Exhibit 1 if you'd 5 MS. HARTLEY: You have it? That would be 6 good. 7 MR. SHEFFLER: We'll do that. Exhibit 1 8 will be the curriculum vitae of Dr. Sanford 9 Barksy. 10 MS. HARTLEY: Okay. 11 (Plaintiffs' Exhibit 1 was marked for 12 identification.) 13 BY MS. HARTLEY: On that C.V., you have a section 14 Ο. 15 called "publications"; is that correct? 16 A. Yes. 17 And you have them all numbered; is Q. 18 that correct? 19 Α. Yes. Q. Can you tell me what the last number 2.0 21 in your most current C.V. of publications is? 22 A. The last number in the publications 23 section is a 111, but I want to say -- I want to 24 point out something. In the past -- and don't 25 remember when this changed -- I listed my patents 00012 1 together with my publications, and a year or so 2 beforehand, I decided I was going to take them out 3 and list them separately. So the numbers could 4 change as a result of that reordering. But right 5 now, the publications as listed list to 111. 6 Now, the most recent numbers, some of 7 those are just communicated manuscripts which I 8 list under publications but I say they are just 9 communicated manuscripts. 10 Q. What else have you brought today with 11 you to the deposition? 12 A. I brought some notes that I took when 13 I read through the history, and I also wrote a list 14 of my pathology findings. 15 Q. Can you please have the court reporter 16 mark that as Exhibit No. 2. MR. SHEFFLER: Stephanie, there wasn't a 17 18 document request in this case. I'm going to let 19 these be marked, but I do want to say that by doing 20 so I'm not waiving my right to object to further 21 requests for documents et, cetera, in depositions. 22 I mean, obviously the proper course is to serve the

```
23 document request prior to the deposition.
24 Since in this case he has the
25 documents with him and you have requested them and
00013
1 because there doesn't seem to be any prejudice, as
2 long as it's understood that we're not adopting
 3 this as a procedure in the future, I will not press
 4 my objection, and I will allow them to be marked.
          MS. HARTLEY: It's my understanding that
 5
 6 what he brings to the deposition I'm entitled to
7 see and have marked.
         MR. SHEFFLER: Well, we differ.
8
9
         MS. HARTLEY: Court reporter, could you
10 please mark those as Exhibit No. 2.
         THE WITNESS: There are two pages. One is
11
12 the history and one is the path findings. Did you
13 want them both marked as 2?
         MS. HARTLEY: That's fine.
14
15
          (Plaintiffs' Exhibit 2 was marked for
16
          identification.)
17 BY MS. HARTLEY:
18
         Q. What else have you brought with you to
19 the deposition?
20
         A. There are some photographs of the
21 slides that I took with my digital camera.
        Q. Photomicrographs?
23
               Yes.
         Α.
24
               Is it possible for you to get color
          Q.
25 copies of those for the court reporter?
00014
1
         MR. SHEFFLER: We do have color copies.
         MS. HARTLEY: Okay. Can you please mark
2
3 those as Exhibit No. 3.
         MR. SHEFFLER: We'll take that under
 5 advisement. I'll let you know at the end of the
 6 deposition.
 7 BY MS. HARTLEY:
8
         Q. How many photomicrographs do you have,
9 Dr. Barksy?
        A. I believe it's ten.
Q. And you took those, yourself, with
10
11
12 your digital camera?
         A. Yes, and we have actually files of
13
14 those, too. Electronic files. If you'd like.
        Q. What else did you bring, Dr. Barksy?
15
16
               I have one of my articles. I have a
          Α.
17 reprint of one of those articles.
18 Q. Which article is it?
19
          Α.
               It's my paper on the rising incidence
20 of bronchioloalveolar lung cancer that was
21 published in "Cancer" in 1994.
22
          Q. What else did you bring with you?
23
          A. Just a tablet to write on if I have
24 to.
25
          Q.
              A blank one?
00015
               Yes.
1
          Α.
 2
               Have you written any notes on it so
          Q.
 3 far?
 4
               No. It's just a clipboard with
          Α.
 5 papers. They are blank.
 6 Q. Now my understanding is that in the
 7 summer of 1999 Bruce Sheffler brought you some
```

```
8 slides to review; is that correct?
9
         A. Correct.
10
          Q.
               What slides did you review? Can you
11 identify them for me by number?
12
          A. No, because I reviewed them in, like I
13 said, in the summer of 1999 and then shortly
14 thereafter I took pictures of them. But I did not
   code the pictures in terms of the numbers of the
15
16 slides. But I recall it was around slightly over
17 ten slides, but I haven't reviewed them recently,
18 and I haven't looked at them recently, and I don't
19 know their numbers, per se.
               Are those these slides that you took,
20
          Q.
21 the photomicrographs?
22
          Α.
               Yes.
23
               And did you take them at that time in
          Ο.
24 the summer of 1999?
2.5
         Α.
               Yes.
00016
1
               Do you know from recollection or from
 2 looking at those pictures whether they were slides
   from the lobectomy or slides from the bronchial
 4 brushing?
 5
          A.
                I recall reviewing the slides of the
 6 bronchial biopsy, but the pictures that I took are
7 from the lobectomy specimen.
8
          MR. SHEFFLER: Stephanie, if you are just
9 looking for information here, I can give the
10 numbers of the slides that he reviewed if you are
11 interested in that.
          MS. HARTLEY: I am.
12
13
          MR. SHEFFLER: I will get those for you.
14 Just leave this area, if you would. I'll get those
15 numbers for you, and I'll read them into the
16 record.
          MS. HARTLEY: Okay. I may some other
17
18 questions.
19
          MR. SHEFFLER: Sure, absolutely. But I mean
20 instead of just fumbling around, I can read you the
21 slides.
          MS. HARTLEY: That will be fine.
2.2
               Did you look at any bronchial brushing
2.3
          Q.
24 pathology?
25
               I don't recall looking at those.
00017
1
          MR. SHEFFLER: In fact, here are the
 2 slides. Okay. Here are all the slides that we've
 3 got. And they are 12018-96V recut; 12496-96FSA-1V;
 4 12496-96FSB-1; 12496-96FSC1; 12496-96FSC2;
 5 12496-96A-2; 12496-96BMHC1; 12496-96B-2;
   12496-96C3. C4, C5, C6, C7, C8, C9 all in the
 6
 7
   series 124-96-96.
                The next series is 8390-98. These are
8
9 all recuts and they are numbered sequentially 1, 2,
10 3, 4, 5, 6, 7. Another 7V. And then there is a
11 slide 15707-98. There's a slide 47962-98MH.
12
                There's a slide 67 -- excuse me.
13 Strike that.
                Slide 6117-99AW; 6117-99BW;
14
15 10916-991W; 10916-992W and 10916-993W. That's it.
16
          MS. HARTLEY: Okay. Thank you.
17
          Q. Dr. Barksy, when you reviewed these
18 slides in the summer of 1999, you told me you
```

```
19 reviewed the slides but you did not have the
20 medical records; is that correct?
21
         A. That's correct.
22
              Did you have any of the pathology
         Q.
23 reports?
24
              I may have had a copy of the path
25 record, but I don't remember for sure. But the
00018
1 medical records followed shortly thereafter in
 2 which I reviewed them and they'd certainly have the
3 path report in them.
         Q. Correct. Did you have anything else
4
5 when you reviewed those slides?
         MR. SHEFFLER: With respect to the case
6
7 specifically?
8
         MS. HARTLEY: Right, or textbooks or
9 anything.
10
              When you were reviewing those slides,
        Ο.
11 what did you have with you?
12
         MR. SHEFFLER: I object to the question.
13 It's overbroad unless you limit it to what did he
14 review when he reviewed the slides. Obviously, his
15 entire library.
16
         MS. HARTLEY: I'm not talking about what did
17 he have in the room with him. I'm talking about
18 when he reviewed the slides.
19
         Q. Dr. Barksy when you reviewed the
20 slides in the summer of '99 in the Boerner case,
21 what did you review at the time you reviewed the
22 slides?
23
              Nothing, I just reviewed the slides.
         Α.
24
               Did Bruce Sheffler give you any
         Q.
25 history on the case at that time?
00019
               Well, when he came to visit me, he did
1
 2 tell me -- he did summarize the case very briefly.
   I mean, he told me it was a woman. I think he told
 4 me her age. I think he told me, you know, a
 5 summary of the history as he knew it.
 6
    Q. Did he tell you that she was a
 7 smoker?
8
               I think he told me that she was a
9 former smoker or an ex-smoker. I think he said
10 that to me at that time.
11
    Q. Did he tell you how many pack years
12 she had smoked?
13
        A. No.
14
         Q. Did he tell you her occupation?
15
              No.
         Α.
16
         Q. Can you recall what summary he gave
17 you of the case?
18 MR. SHEFFLER: Objection. Asked and
19 answered, but go ahead.
20 THE WITNESS: Well, in addition to the
21 things I already mentioned, I think he mentioned to
22 me that this was a case that was going to go to
23 trial. That it was in Arkansas. It was in federal
   court. That's what I remember he said about it.
24
25 BY MS. HARTLEY:
00020
               Did you take the pictures at that
1
 2 meeting?
 3
         A. Well, what happened was he left the
```

```
4 slides with me and I took them that evening and the
 5 next day he picked them up.
 6
          Q.
              So did you take photomicrographs that
7 evening?
8
         Α.
               Yes.
9
               And those are the ones you have with
          Q.
10 you today?.
11
          Α.
               Yes.
12
         Q.
               So the medical records followed?
13
         Α.
               Yes.
14
              Did he send you anything else aside
15 from the medical records?
16
         Α.
               No.
               When he sent the medical records, did
17
          Q.
18 he come and meet with you to discuss them?
19
         A. No.
20
               Did you discuss them with him over the
         Q.
21 phone?
22
              No.
23
               Let me modify that. I didn't discuss
24 the medical records, per se, with him. When they
25 came, I reviewed them, but after that, a few weeks
00021
1 or a month or so after that, Mr. Sheffler did call
 2 me and we discussed my whole impression of the
4
               Have you reviewed any radiology in
          Ο.
 5 this case?
 6
         Α.
               No.
 7
          Ο.
               Have you consulted with any other
8 doctors in this case?
9
         A. No.
10
         Q. Have you discussed the case with
11 anyone other than Mr. Sheffler or people in his
12 office?
13
               No.
          MR. SHEFFLER: Let me just -- Dr. Barksy may
14
15 not know. Dr. Woodside did have a brief discussion
16 with Dr. Barksy about this case. He's here today.
17
         MR. WOODSIDE: Only in your presence.
18
         MR. SHEFFLER: Right, with me so I just want
19 the record to reflect.
20 BY MS. HARTLEY:
21
          Q.
              Did you review Dr. Feingold's
22 deposition in this case?
23
         A. No.
         Q. Have you seen Dr. Feingold's report?
24
25
         Α.
              No.
00022
1
         Q. Did you review Dr. Sidransky's
 2 deposition in this case?
          A. No.
 3
4
          Q.
               Have you reviewed Dr. Sidransky's
 5 report?
 6
              No.
          Α.
7
               Have you seen photomicrographs taken
          Q.
8 by anybody else in this case?
             No.
         A.
9
               Have you been told about
10
          Ο.
11 Dr. Feingold's testimony in this case?
          A. The only thing I recollect is somebody
12
13 mentioned -- I guess it was Mr. Sheffler -- that
14 Dr. Feingold was involved in the case, but I have
```

15 not been told anything about his testimony. Q. Have you been told anything about his 16 17 impression? 18 Α. No. 19 Q. Have you been told anything about 20 Dr. Sidransky? 21 Α. No. 22 Do you know who Dr. Sidransky is? Q. 23 Α. Yes. 2.4 What is your understanding of who Ο. 25 Dr. Sidransky is? 00023 I believe Dr. Sidransky is a head and 1 2 neck physician who's also a scientist and does 3 cancer research. I believe he's at Johns Hopkins. 4 Q. Has anyone told you of Dr. Sidransky's 5 impressions in this case? 6 A. No. 7 Have you read any depositions at all Q. 8 regarding this case? 9 Α. No. Dr. Barksy, since I have the benefit 10 Q. 11 of your current C.V. in front of me, can you tell 12 me whether you've written on lung cancer in the 13 past five years? 14 Α. 15 Ο. And which articles would that be? 16 I've written some abstracts, and I've Α. 17 also written a paper that's going to be published, 18 and I have another paper that's in the -- it's 19 about to be communicated. 20 Can you tell me what numbers those are Q. 21 on your C.V.? 22 Α. Sure. Let me just check it out. This is under a heading that's called 2.3 "Abstracts and Presentations." Number 64 is an 24 25 abstract by authors O'Connell, Heras, Palmarini, 00024 1 Sharp and Barksy that's entitled "JSRV-related 2 Sequence and Capsid Protein in Human Lung BAC/PAC 3 Suggests a Retroviral Connection." It's published 4 in "Lab Investigation" in 1998. The next one under "Abstracts and 5 6 Presentations" is number 75. It's by Grossman, Hiti, McNiel, O'Connell, Shao and Barksy entitled 7 "Feline Bronchioloalveolar Lung Cancer Shares 8 9 Common Properties with Sheep and Human BAC." It's 10 in "Experimental Biology" 1999. 11 The next is number 84. It's just by 12 Barksy. It's entitled "A Retroviral Link to Human Tobacco-Related Lung Cancer?" And it was in what 13 14 was called the "AIM99: Tobacco Research in Action 15 Symposium" in 1999. 16 Q. What was the title of that? 17 "A Retroviral Link to Human 18 Tobacco-Related Lung Cancer?" That's it for the 19 abstracts. 20 Now, in the column that's entitled 21 "Publications," there are a number of publications 22 on lung cancer. The first is number 72. The authors 23 24 are Fligiel, Roth, Kleerup, Barksy, Simmons and 25 Tashkin entitled "Tracheobronchial Histopathology

00025 1 in Habitual Smokers of Cocaine, Marijuana and/or 2 Tobacco, "published in "Chest" volume 112:319 to 3 326, 1997. The second paper is numbered 79 by 5 Roth, Arora, Barksy, Kleerup, Simmons and Tashkin entitled "Airway Inflammation in Young Marijuana 6 7 and Tobacco Smokers" in the "American Journal of 8 Respiratory Critical Care Medicine, "volume 157;928 9 to 937, 1998. 10 Next is paper No. 85 by authors 11 Barksy, Roth, Kleerup, Simmons and Tashkin entitled "Histopathologic and Molecular Alterations in 12 13 Bronchial Epithelium in Habitual Smokers of 14 Marijuana, Cocaine and/or Tobacco" published in the 15 "Journal of American Cancer Institute" volume 16 90:1198 to 1205, 1998. The next is number 99 by De las Hera, 17 18 Barksy, Hasleton, Wagner, Larson, Egan, Ortin, 19 Gimenez, Palmarinin and Sharp entitled "Evidence 20 for a Protein Related Immunologically to the 21 Jaagsiekte Sheep Retrovirus in Some Human Lung Tumors." This is in press in the "European 22 23 Respiratory Journal." 24 The next is number 110 O'Connell, 25 De las Heras, Palmarini, Sharp and Barksy. Title 00026 1 is "JSRV-related Sequence and Capsid Protein in 2 Human Lung BAC/PAC Suggests a Retroviral 3 Connection" and this paper has been communicated to 4 the "Journal of Virology," but it has not been 5 accepted yet. 6 There's one other number 106 and it's 7 by Grossman, O'Connell, Hackett, McNiel, Hiti, 8 De las Heras, Sharp and Barsky. It's entitled "Feline Bronchiloalveolar Lung Carcinoma and a 9 10 Derived Cell Line Share Common Properties with Both 11 Sheep Pulmonic Adenomatosis and Human BAC." And 12 actually, there's an error here. It is submitted 13 to "Cancer Research," but it's not yet accepted. I 14 had written that it was in press, but that's not 15 correct. That's an error. It's a submitted 16 stage. 17 That's it. Oh, there's one other thing. There's 18 19 one other item. This is under my U.S. patents. 20 It's by Barksy and Grossman. A feline 21 bronchioloalveolar lung cancer xenograft and cell 22 line for the study of common animal-human 23 pathogens. It's an United States patent 24 application. That's it. 25 Do you get funding for any of these Ο. 00027 1 articles that you've written? Α. 3 Who did you get funding from? Who 4 have you gotten funding from in the last three 5 years? MR. SHEFFLER: Objection. For these 6 7 articles or for everything? 8 MS. HARTLEY: For these articles. 9 THE WITNESS: Well, there are different 10 funding sources for different articles. Some of

- 11 the funding comes from the National Institutes of
- 12 Health. Some comes from a private foundation in
- 13 Los Angeles called the Margaret Early Foundation,
- 14 and some comes from the UC, University of
- 15 California Tobacco Related Disease Research
- 16 Program.
- 17 BY MS. HARTLEY:
- 18 Q. Does any of the funding come directly
- 19 from the tobacco companies?
- 20 A. No.
- Q. When you reviewed the slides back in
- 22 the summer of 1999, can you tell me what your
- 23 impressions were?
- 24 A. Let me first sort of clarify or
- $25\,$  embellish something. I said that I reviewed the  $00028\,$
- 1 slides in the summer of '99 and then I took digital
- 2 pictures of what I thought the most cellulin areas
- 3 were. I have reviewed not the slides again but I
- 4 have re-reviewed those pictures, and I've reviewed
- 5 those pictures, you know, a number of times and
- 6 even recently. So when you ask me what my opinions
- $7\,$  were in the summer 1999, you know, it's hard to
- 8 really dissect that out from what my opinions are
- 9 currently. Not necessarily that they've changed,
- 10 but it's hard for me to think back of my thought
- 11 processes at that time. I just wanted to clarify
- 12 that point.
- Q. Well, if you cannot differentiate
- 14 between what you thought then and what you think
- 15 now, then that's fine.
- 16 A. I just wanted to clarify that with
- 17 you.

6

- 18 Q. Is it correct, Dr. Barksy, you cannot
- 19 separate out what you thought when you first
- 20 reviewed the slides with what you now think that
- 21 you have reviewed the slides and the medical
- 22 records and had other information provided to you?
- 23 A. Well, I don't feel my opinions have
- 24 changed, but on the other hand, you know, what I'm
- 25 thinking now is what I'm thinking now, and I just 00029
  - 1 wanted to make that point to you.
  - Q. So is that correct, Dr. Barksy, that
  - 3 you cannot differentiate for me what you believed
    4 in the summer of 1999 when you first saw the slides
  - 5 and what you think now?
    - A. Well, I can do the best I can.
  - 7 Q. Okay. Well, then can you tell me what
- 8 you believed when you reviewed the slides in the 9 summer of 1999?
- 10 A. Okay. When I reviewed the slides --
- 11 and like I said I have reviewed approximately 10 or
- 12 more slides of this mass lesion in the left upper
- 13 lobe of the lung -- I saw a number of different
- 14 histopathological patterns. The mass was clearly a
- 15 primary lung cancer. I saw areas of
- 16 bronchioloalveolar lung cancer. And these areas
- 17 were of the nonmucinous type. They seemed to be of
- 18 the Clara cell type, and they were lined juxtaposed
- 19 to a pulmonary scar, which was filled with a lot of
- 20 elastosis. And the scar suggested to me that it
- 21 was due to an old pulmonary infarct of the lung

22 because of the presence of elastosis. 23 Also around the scar, I found foci of 24 what I would term bronchioloalveolar metaplasia, 25 hyperplasia and atypical hyperplasia. These foci 00030 1 progressed into the areas of bronchioloalveolar 2 lung cancer. Now, in addition, there were other 3 4 areas of this tumor that were much poorly 5 differentiated, and they were what I would consider 6 areas of dedifferentiation, and these areas were 7 varied. Some were adenocarcinoma. Some were 8 squamous cell carcinoma. Some were a mixture of 9 adeno and squamous, which I would term 10 adenosquamous carcinoma. There were areas that 11 were differentiated and which I would term Clara 12 cell carcinoma, but many of these areas were kind 13 of mixed together. So the way I put the case together in 15 the lung, that this was a scar-related 16 bronchioloaveolar lung cancer that was arising from 17 these foci of metaplasia and hyperplasia and 18 atypical hyperplasia, and that it had progressed 19 and dedifferentiated into aggressive appearing 20 adenocarcinoma and squamous cell carcinoma and 21 adenosquamous carcinoma and Clara cell carcinoma. 22 I found a small metastasis in one of 23 the bronchial lymph nodes, and I also recall seeing 24 that the bronchus, the tumor was fairly large. It 2.5 filled most parts of the lung. It grew between the 00031 1 pleura. It was subpleural in areas next to the 2 scar, but it did extend to grow next to a major 3 bronchus, but the bronchus was histologically 4 normal in appearance. And that's what I remember seeing in 5 the summer, and as I reviewed the pictures that I 6 7 took, my opinion was confirmed and reinforced. But 8 my opinion, you know, right now is what I have 9 today, and what I'm telling you. That was just the 10 point I wanted to make. So your opinion today is what you just 11 Q. 12 told me? 13 Yes. 14 Ο. You said -- first, let me go over 15 this. I have made some notes. I'd like you, if 16 you would, to pull out your pictures. Are those 17 pictures numbered? 18 Yes, well they have letters. Α. 19 They have letters? Q. 20 Α. Yes. 21 Well, what I'd like to do is as I talk Ο. 22 about these areas that you saw, I'd like you to 23 tell me which picture best represents that. I 24 believe we've done this before Dr. Barksy. 25 You said that you saw areas of 00032 1 nonmucinous BAC; is that correct? 2 Α. Yes. 3 Which pictures best represent that Q. 4 nonmucinous BAC? MR. SHEFFLER: We're going to take a short 6 break here to consult, Stephanie. We'll be right

```
7 back.
 8
          MS. HARTLEY: Okay.
9
          (A brief recess was taken.)
10
          MR. SHEFFLER: What we're going to do --
11 since you want to talk about the pictures, let me
12 tell you what we have. We've got a code for the
13 pictures that Dr. Barksy has photographed. It has
   the ten pictures and their description, and we have
14
15 the pictures themselves. Now, you want these
16 marked or how do you want to proceed?
17
          MS. HARTLEY: Yes, I would like to have them
18 marked.
          MR. SHEFFLER: May I make a suggestion? May
19
20 \, I suggest that you mark the code as 3 and then the
21 pictures as 3-A through -J because they are already
22 labeled A through J.
         MS. HARTLEY: That's fine.
23
          MR. SHEFFLER: Exhibit 3 and A through J.
2.4
          (Plaintiffs' Exhibits 3, 3-A through 3-J
00033
          were marked for identification.)
1
 2 BY MS. HARTLEY:
               Dr. Barksy, before we took a break, I
 3
          Q.
 4 had asked you if you would identify for me the
 5 picture that best represents the nonmucinous area
 6 of BAC that you had described previously.
 7
               Well, in the pictures I took, the
8 nonmucinous BAC is present on at least three of the
   pictures, and they all show the BAC very well. So
9
10
   I don't know which the best one is.
11
          Q. Why don't you tell me all three?
12
               Well, in the slide that's labeled
          Α.
13 Boerner A. That's one of them. Then there is a
14 slide labeled Boerner F, which has a scar with the
15 BAC. And then there's a slide labeled Boerner J,
16 which is also BAC.
17
          Q. And those are the slides that you
18 believe represent nonmucinous areas of BAC?
          A. Correct.
19
20
               Then you said it was juxtaposed to a
21 pulmonary scar and you told me that slide F shows
22 that scar; is that correct?
              Slide F, yes.
23
          Α.
               Are there any others that you believe
24
          Q.
25 demonstrate the pulmonary scar that you have seen?
00034
1
               Yes. The foul F or slide F was the
 2 scar with BAC, but we have a picture just of the
 3 scar, and that's labeled Boerner E.
 4
               Any others?
          Q.
 5
          Α.
                No.
 6
                Then you said that there was an area
          Ο.
 7
   of adenocarcinoma?
8
          Α.
               Yes.
9
          Q.
               What appeared to you to be
10 adenocarcinoma?
11
          Α.
               Yes.
12
                What slides best represent that?
          Q.
13
               Well, there's a slide that has adeno
          Α.
14 and squamous carcinoma on the same slide and even
15 also has some Clara cells and so that's Boerner B.
16
          Q. Any others?
17
          Α.
               No. No other slides or any others
```

18 that represents that squamous, Clara cell. 19 Q. Any other slides that represent some 20 adenocarcinoma? 21 Α. 22 Q. Any other slides that represent some 23 squamous carcinoma? A. No. I tried to photograph -- I used 24 25 this slide because it -- I saw areas that were 00035 1 squamous and adeno and that, but I didn't 2 photograph them. I photographed those areas on 3 this slide alone. Q. Okay. Now how about areas of 4 5 adenosquamous carcinoma? Which slide best 6 represents that? 7 A. Well, that would also be in the slide 8 в. 9 And you said you saw some Clara cell 10 carcinoma. Any other slide besides slide B that 11 represent that? 12 There are other areas, but I didn't Α. 13 photograph them. Q. So we have now slide A, B, E, F and 14 15 J. 16 What does slide C represent? 17 Slide C is a section of a normal 18 bronchus. A normal appearing bronchus. And slide D? 19 Q. It's a normal appearing bronchus that 20 Α. 21 has adjacent tumor. 22 Q. And slide G? That's a focus of bronchioloalveolar 23 Α. 24 metaplasia. And slide H? 25 Q. 00036 A. That's a focus of bronchioloalveolar 1 2 hyperplasia. And slide I? Q. 3 That's a focus of bronchioloalveolar 4 Α. 5 hyperplasia with atypical cells. Q. So if asked what type of cancer you 6 7 believed Mary Jane Boerner had, what would your 8 answer be? 9 I would say it was a 10 bronchioloalveolar lung cancer that had 11 dedifferentiated into adenosquamous carcinoma, 12 adenocarcinoma, squamous cell carcinoma, and Clara 13 cell carcinoma. That would be my "official 14 diagnosis" if it was a case that I was reviewing in 15 my routine duties as a pathologist. Do you have an opinion as to what 16 Q. 17 caused Mary Jane Boerner's lung cancer? 18 A. No. 19 Do you have an opinion as to what risk 20 factors may have contributed to Mary Jane Boerner's 21 lung cancer? I think that the scar which is present 22 23 may have been a risk factor. The scar from an old 24 infarct. 25 Anything else? Q. 00037 1 Α. 2 Q. Do you have any opinion on what may

3 have caused the scar? 4 A. I think the scar was due to an old 5 infarct, and that's due to either poor circulation 6 or a thromboembolus. They are fairly common in the 7 lung in patients you know, of her age. Q. Is smoking a risk factor for poor 8 9 circulation? 10 A. I'd say in general, yes. It's usually 11 arterial circulation, though, to the extremities or 12 to the heart. 13 Q. Is smoking a risk factor for 14 thromboembolism? 15 Α. Yes. Are you familiar with the WHO 16 Q. 17 classifications of tumors? 18 A. Yes. 19 Q. Do you consider the WHO 20 classifications of tumors as represented in their 21 treatise of the Armed Forces Institute of Pathology 22 to be authoritative? MR. SHEFFLER: Object. Do you really want 23 24 to say the WHO in their treatise AFIP? MS. HARTLEY: Right. 25 00038 1 MR. SHEFFLER: I object. Those are two 2 different organizations, and that's non sequitur. 3 MS. HARTLEY: Let me ask it again. Q. Dr. Barksy, do you believe that the 4 5 WHO classification of tumors as reflected in their 6 treatise is authoritative? 7 A. Yes, I do. But let me just qualify my 8 answer by saying it's not the only agency or group 9 of pathologists that make criteria. There are 10 other bodies. There are other treatises. There 11 are other textbooks. There are other societies 12 that come up with criteria as well. Those are 13 also -- I consider those also authoritative. In addition, in pathology, there are 14 15 the authoritative nature of learned scholars, 16 authors of textbooks, et cetera. One takes not any 17 single agency or group and believes only what they 18 have to say exclusively. One has to look at the 19 whole -- the whole field and then one adopts, you 20 know, what the consensus is and one applies one's 21 own common sense out of the whole, you know, 22 milieu. So although I do consider them 23 24 authoritative, I don't rely solely on what the WHO 25 declares as being truth. I have to look at the 00039 1 whole ball of wax, if you will. 2 Q. What did you rely on in terms of 3 treatises in concluding that this was a BAC lung 4 cancer that had dedifferentiated into adeno, 5 squamous and Clara cell? MR. SHEFFLER: Objection to the form. 6 7 Assumes facts not in evidence. In fact, assumes 8 facts contraindicated by the previous testimony. 9 BY MS. HARTLEY: 10 Q. Go ahead and answer. 11 Well, when I look at a case, any case, 12 I bring to the microscope the sum total of my 13 training and experience. That means what I've

14 read, what I've seen. I formed over the years as a 15 pathologist my own criteria. If I had to go back 16 to the books on every case that I look at, I would 17 never get through my daily workload. So this case is no different. I 18 19 brought the sum total of my experience, my own 20 published work, the work of other organizations 21 such as the WHO et, cetera, and applied to the best 22 of my ability an interpretation of the patterns 23 that I saw before me. Dr. Barksy, does smoking cause lung 24 25 cancer? 00040 It depends on what you mean by 1 "cause." Certainly in populations for certain 2 3 cancers of the lung it contributes a tremendous 4 increased risk. And from an epidemiological 5 standpoint, it causes lung cancer. But again, it 6 depends what you mean by "cause." If you speak of 7 an individual case, it may cause an individual case 8 of lung cancer, but you have to apply different 9 criteria to causation than you do 10 epidemiologically. 11 Do you believe smoking caused Mary Q. 12 Jane Boerner's lung cancer? 13 A. No, I do not. 14 And why do you not believe her smoking Ο. 15 caused her lung cancer? A. Well, the main reason is because I 16 17 feel the cancer that she developed was a 18 bronchioloalveolar lung cancer and it seems to be 19 arising in a preexisting scar. That's the first 20 reason. The second reason is I see evidence 21 22 under the microscope of a progression from 23 precursor lesions of bronchioloalveolar metaplasia and hyperplasia, which are lesions that exist in 24 25 the periphery of the lung and which are thought to 00041 1 be precursor lesions of bronchioloalveolar lung 2 cancer. The third component of the evidence as 3 4 I put the case together is that this patient wasn't 5 an active smoker but was an ex-smoker, who I 6 believe from reviewing the chart has stopped for 7 over 15 years. So she's in the category of 8 ex-smoker rather than active smoker. And although 9 ex-smokers have an increased rate of lung cancer, 10 the types that they get and their overall incidents 11 decrease progressively with years of secession. 12 So all these things are in my mind 13 when I come to my conclusion. But it's mainly 14 because of the histological type, the fact that I 15 believe it is a BAC, and that it's arising 16 peripherally from these precursor lesions. And 17 finally, I don't see any evidence of a central 18 airway abnormality in this patient. 19 Q. Have you ever reviewed a case that you 20 believed -- a case of lung cancer that you believe 21 was caused by smoking? 22 A. Oh, sure. 23 Q. How many? 24 Well, you have to understand that in Α.

00042 1 cancer, I'm not really asked to address that 2 issue. I'm not asked to address etiological 3 concerns. I'm simply asked to I diagnose and 4 categorize the particular kind of lung cancer. you know, I'm not routinely thinking on those 5 6 7 But when I've been asked to look at 8 cases by attorneys for cigarette companies, 9 etcetera, I have reviewed cases that I have felt 10 that it was likely that cigarette smoking played a 11 role in the causation or etiology of the particular 12 case. 13 And can you recall the names of any of Ο. 14 those? 15 MR. SHEFFLER: Objection. I think it calls 16 for work product. I'm going to instruct the 17 witness not to answer. 18 I just instructed you, Dr. Barsky, not 19 to answer. You may ignore my instruction if you 20 wish, but that may be work product, and I'm not 21 sure she's entitled to that. 22 BY MS. HARTLEY: 23 Q. Does cigarette smoking cause BAC? 24 BAC is a type of lung cancer in which 25 I've not been convinced that there is convincing 00043 1 epidemiological evidence in terms of odds ratio and 2 relative risk that meet the criteria 3 epidemiologically of causation. There are some 4 studies that show a weak association but not strong 5 enough to implicate causality. I never think of 6 BAC or I haven't thought of BAC as one of the 7 cancers of the lung that are linked to smoking. What do you associate BAC with in 8 Q. 9 terms of causation? 10 A. I think that the vast majority of 11 cases of BAC the causation is unknown. We know 12 we're seeing an increase in BAC and also in 13 peripheral adeno. We're seeing an increase in 14 patients who smoke as well as who don't smoke, and 15 it's not clear why that's occurring. 16 There have been a number of risk 17 factors over the years implicated. A high fat 18 diet, perhaps radon exposure. There's is a viral 19 hypothesis that's resurrected from time to time, 20 which I'm actively studying right now, but I think 21 most BACs occur and certainly a large number do 22 occur in nonsmokers and there's no apparent 23 etiological agent that one can implicate. 24 Do you know if Mary Jane Boerner had a Q. 25 high fat diet? 00044 1 I recall a statement in her chart that 2 said she was obese. I assume that meant she had a 3 high fat diet, but doesn't mean she did, per se. 4 All you know about her diet was a Q. 5 mention in the chart that she was obese? 6 She had a problem, I remember reading Α. 7 in the chart after her cancer was diagnosed where 8 she was having thromboembolic phenomena especially 9 in her left hand and arm. And one of the

25 my routine job, when I review cases of lung

```
10 therapists in addition to being anticoagulated, I
11 remember something to the effect that they wanted
12 her on a low cholesterol diet. I assume that meant
13 her cholesterol was elevated, but I don't recall
14 specifically seeing that. It may have been there,
15 but I don't recall reading that.
               From seeing her cholesterol level?
16
          Q.
              Yes. I'm sure they are there, but I
17
          Α.
18 just don't recall. I don't recall what those
19 were.
          Q. Do you have any knowledge on whether
20
21 Mary Jane Boerner was exposed to radon?
2.2
         Α.
               No.
23
               Do you have any knowledge on whether
          Q.
24 Mary Jane Boerner had any kind of virus that may
25 have caused -- virus that you associate as causing
00045
1 BAC?
          Α.
               No.
3
          Q. Do you believe that the risk of
4\, developing lung cancer returns to 0 after a person
5
   stops smoking?
          MR. SHEFFLER: Object to the form of the
6
7 question. It makes no sense if you talk about
8 relative risk. There is no 0 relative risk.
9
          THE WITNESS: There are a number of
10 studies. Some are controversial, some studies, or
11 some with conflicting result. There are studies
12 that show that one's risk for lung cancer does
13
   decrease with the number of years of stopping. And
14 that it approaches the risk of the nonsmoking
15 population after so many years.
16
               The studies are different though in
17 terms of the number of years. Some say that you
18 approach a baseline risk about 15 years after
19 you've stopped. Some say at 25 or 30 years. Some
20
   say you never really approach the normal risk of --
21 the nonsmoker's risk of developing lung cancer.
22 But most studies agree that the risk does decrease
23 with the number of years. I mean, that's the whole
24 basis of urging people to stop smoking cigarettes.
25
                But the studies are, you know, a bit
00046
1 controversial, and I don't have a hard fast
 2 opinion. I believe the risk does decrease.
 3 still think it's present, but I don't know exactly
 4 at what point or if it ever gets exactly to what
 5 the nonsmoker risk is. But what I do know is that
 6 there are many etiological factors out there,
 7 especially for BAC, that are not defined that the
 8
   vast majority of patients with BAC do not have a
9 risk factor but they get the tumor anyway.
10 BY MS. HARTLEY:
11
          Q.
               Did you perform any tests on the
12 pathology material?
13
          Α.
               No.
               Do you know of any tests that were
14
          Q.
15 performed on the pathology material?
16
          A. No.
          Q. Are you board certified, Dr. Barksy?
17
               Yes. In anatomic and clinical
18
          Α.
19 pathology.
20
          Q. You're not a pulmonary pathologist,
```

```
21 are you?
22
                I am because I'm in charge of
          Α.
23 reviewing all the pulmonary pathology at UCLA,
24 especially the lung cancer cases. I don't have
25 subspecialty boards in pulmonary pathology, and I
00047
1 don't even know if they have subspecialty boards in
   that particular specialty, but I have become
 3 through my publications and through my job
 4 performance a pulmonary pathologist, especially for
 5 neoplastic pulmonary diseases. Not for
 6 non-neoplastic diseases. That's a separate kind of
 7 specialty.
                Have you ever read the Surgeon
8
          Q.
9
   General's report relating to smoking?
10
          Α.
               I've never actually read the Surgeon
11 General'S report, but I've certainly seen them
12 quoting in text, newspapers and other journal
13 articles.
14
          Q.
                Would you recognize them as
15 authoritative?
          MR. SHEFFLER: Objection to the form of the
16
17 question, and also object that it's overbroad.
18 Authoritative as to what? Which reports and to
19 what subjects?
20
          THE WITNESS: I would consider them, you
21 know, somewhat authoritative. They are actually
22 based on epidemiological studies that preceded the
23 reports by five to ten years.
24 BY MS. HARTLEY:
25
                Could you hold on, Dr. Barksy. I
          Q.
00048
1 think I'm almost finished.
                Dr. Barksy, have you reviewed any
 3 tobacco company documents? And what I mean by that
 4 is documents that were offered by the tobacco
 5
   companies that have not been published.
 6
           Α.
                No.
 7
                Have you read any of the recent books
           Ο.
 8
  that have been out for the past six or seven years
 9 on the tobacco company and cigarette smoking?
10
          Α.
               No.
          MS. HARTLEY: I believe those are all the
11
12 questions I have, Dr. Barksy.
13
          MR. SCHULZE: This is Gerry Schulze, and
14 ordinarily I don't like to double team, but I was
15 provided with a curriculum vitae for Dr. Barksy
16 yesterday, and I'm just going to ask Stephanie
17 while we're on the record because I think everybody
18 is going to know where I'm going. It contains a
19
   summary of areas of Dr. Barksy's interest, I
20 suppose, and in that, Dr. Barksy states that BAC is
21 a form of lung cancer who's etiology and
22 pathogenesis is controversial and whose link to
23 either main stream tobacco smoking or secondhand
24 smoking unproven, and then he goes on to say past
25 studies by the P.I. suggest that this type of lung
00049
 1 cancer has increased dramatically in the last
 2 decade and is now the most common type of lung
 3 cancer seen at UCLA.
                Stephanie, have you seen that
 5 statement?
```

```
6
          MS. HARTLEY: No.
 7
          MR. SCHULZE: Do you mind if I ask
 8 Dr. Barksy a few questions about that?
9
         MS. HARTLEY: No, I don't mind.
          MR. SHEFFLER: She might not mind, but I
10
11 think I might.
          MR. SCHULZE: Do you mind?
12
13
          MR. SHEFFLER: Go ahead, Gerry.
14
15
                       EXAMINATION
16 BY MR. SCHULZE:
17
          Q. Doctor, you state that BAC is one of
18 the most common types of lung cancer seen at UCLA;
19 is that correct?
20
         Α.
21
          Q.
               What is your source for that? Has
22 somebody done a study?
23 A. I did a study.
               Personally did a study?
          Q.
25
          Α.
               Well, there's two components of that.
00050
1 I review all the lung cancers at UCLA, and over the
 2 years I had noticed, anecdotally at least, that I
 3 was seeing more and more cases of BAC. And some of
 4 my surgical colleagues were telling me that they
 5 were seeing more and more cases of BAC. So I
 6 decided to see if this was really true or it was
 7 just, you know, my own anecdotal bias.
                So the study that I did in 1994, I
8
9
   re-reviewed and pulled all the cases at UCLA from
10 the '50s to the '90s and saw that, in fact, the
11 absolute number and the relative number of cases
12 were increasing. And that increase has -- it's
13 sort of leveled off recently, but it's still the
14 most common type of lung cancer we see at UCLA.
          Q. Are there any published studies that
15
16 you're aware of that any other institution has seen
17 that type of increase in BAC?
18
          A. Yes. There are a number of studies
19 from other institutions that have -- and from the
20 United States as well as other countries that have
21 noted an increase in BAC. Some of these studies
22 have also noted an increase in peripheral
23
   adenocarcinomas. Non BAC peripheral adeno. But I
24 think it's a general consensus that peripheral
25 adenos and BAC have increased in the last decade
00051
1 and even before.
                There are a number of studies.
 3 can't quote them, you know, off the top of my head
 4
   at the moment, but I know that my paper cites some,
 5
   and there have been others that have been observed
 6 in the literature.
 7
          Q. Do you have any theory as to a reason
8 for the increase in BAC?
9
               Obviously, any time you have a disease
10 that has increased in incidence there has to be a
11
   "new factor in on the horizon" to explain it. And
12 that was one of the reasons why my interest in the
13 retroviral etiology was resurrected. Now, that
14 doesn't mean that that's the reason, but it means
15 there is a factor that has emerged or factors,
16 etiological factors that have changed.
```

```
17
                It's just like, you know, when AIDS
18 came on the scene out of nowhere, and there was an
19 AIDS epidemic; that meant that there was something
20 etiological and environmentally that was causing
21 this newly emerging disease and the increased
22 incidence. So it's an intriguing question. It's
23 one of the reasons why I want to study this disease
   and why I have studied it but I don't know for sure
2.4
25 the reason.
00052
               What are the primary types of lung
1
 2 cancer that are recognized?
               Well, the major types are the
 3
          Α.
 4 non-small cell lung cancers which include squamous
 5
   cell carcinoma, adenocarcinoma of the non-BAC type,
 6 the bronchioloalveolar lung cancer, the large cell
 7 undifferentiated cancer, the large cell
 8 neuroendocrine carcinoma, sometimes varying
 9 components of mixed histology. And then of course
10 there are the small cell cancers of the lung which
11 can be subdivided into classic and intermediate and
12 mixed. Those are the major denominations.
          Q. I want to make sure I understand. Are
13
14 you stating here that more cases of lung cancer at
15 UCLA are BAC than any other kind?
16
          A. I think BAC is the most common type
17 that's diagnosed but peripheral adeno, non-BAC is a
18 very close second, and I believe our numbers, they
19 were kind of similar.
20
          Q.
               Have you published the result of that
21 study?
               Well, that's my 1994 study.
22
          Α.
23
          MR. SHEFFLER: Gerry, we'll attach to it the
24 record if you'd like.
          MR. SCHULZE: Okay.
2.5
00053
          MR. SHEFFLER: For the record, it's entitled
1
 2 Rising Incidence of Bronchioloalveolar Lung
 3 Carcinoma and it's Unique Clinicopathologic
 4 Features, and it shall be Exhibit 4.
 5
          (Plaintiffs' Exhibit 4 was marked for
          identification.)
 6
 7 BY MR. SCHULZE:
 8
          Q. I guess what I'm trying to get at, to
9 make sure I understand what the doctor is saying
10 is, are you saying that 51 percent or greater that
11 the lung cancer seen at UCLA is bronchioloalveolar
12 cancer?
13
               No. I said it was the most common
14 type of cancer diagnosed, but that doesn't mean 51
15 percent. It could be 20 percent, 25 percent. As
16 long as it exceeds the other specific types. And I
17 think in this study it was around 25 percent.
18
          MR. SCHULZE: I believe that's all I have.
19 Stephanie, is there anything else?
20
          MR. SHEFFLER: I have a few questions.
21
22
                       EXAMINATION
23 BY MR. SHEFFLER:
2.4
          Q. Dr. Barksy, are you a medical doctor
25 and licensed to practice?
00054
1
          Α.
               Yes.
```

Where are you licensed to practice? 2 Ο. 3 A. Presently California. And if I understand from your answers 4 Q. 5 to prior questions, Dr. Barksy, are you an 6 anatomical and experimental pathologist? 7 Yes. Α. And are you board certified, sir? 8 Q. 9 Α. Yes. 10 And, Doctor, did I understand you to Ο. 11 testify that you are a pulmonary pathologist? 12 A. I consider lung cancer as one of my 13 areas of interest and specialization. And where are you employed? 14 Q. I'm employed at UCLA. 15 Α. At UCLA, Doctor, do you work in the 16 Q. 17 medical hospital? Α. Yes. 18 19 Q. Do you also teach as a professor? 20 Α. Yes. 21 Q. And do you also do research? Α. 22 Yes. 23 Now, in your role as a physician at Q. 24 the UCLA Medical Hospital, do you see lung 25 cancers? 00055 1 Α. Yes. 2 About how many pathologists are at Ο. 3 that hospital? Well, in anatomic pathology, there's 4 5 about ten, but there are other pathologists in 6 clinical and experimental. 7 Q. So there's 10 pathologists. Now is 8 there any pathologist at UCLA who specializes in 9 the diagnosis of lung cancers? 10 Α. Just me. So you are the lung cancer pulmonary 11 Q. 12 pathologist at UCLA? Yes. 13 A. 14 Doctor, could you tell us a little bit Q. 15 about that hospital? A. Well, it's considered a teaching 16 17 hospital. It's considered a quaternary referral 18 center. 19 Q. What does that mean? It means that -- well, there are 20 Α. 21 primary health care centers like at a community 22 level, then there are referral centers perhaps at a 23 municipal level or regional level, and then there 24 are really specialized hospitals that are called 25 tertiary care centers, and then there are the 00056 1 super-specialized hospital, which are called 2 quaternary centers where very specialize things are 3 done and very unusual diseases are treated. And 4 UCLA would fit that latter categorization. 5 Can you give us a rough estimate of 6 how many quarternary specialist hospitals there are 7 in the country? 8 A. There are probably 20, 25, I imagine. 9 And UCLA is one of those 25? Q. 10 Α. Yes. Doctor, could you just briefly -- and Q. 12 I mean very briefly -- tell us what your

13 educational background is starting with med 14 school? Where did you go to med school? A. I went to the University of Pittsburg 15 16 med school. Then I did some training in Internal 17 Medicine at the University of Massachusetts. Then 18 I did my pathology training at Harvard at the Beth 19 Israel Hospital. Then I worked as a pathology for 20 one year at George Washington, and then I did some 21 research training at the National Cancer 22 Institute. 23 Let me just back up for a second. At 24 the National Cancer Institute, Doctor, what type of 25 research was this? 00057 1 Α. I did cancer research specifically on 2 mechanisms of cancer metastasis. 3 Q. Were you still doing your pathology 4 work at the time? 5 Α. 6 Q. After the NCI experience and the 7 research, what did you do? A. Well, then I got my first real 8 academic position where I did research and teaching 9 10 and diagnostic work and that was at UCLA. 11 Q. So at UCLA, Doctor, what is your 12 title? 13 Α. It's now professor of pathology. So are you a full professor? 14 Q. 15 Α. Yes. 16 Ο. And I also noted from your C.V., 17 Doctor, that you are a deputy coroner; is that 18 correct? 19 Yes. Α. 20 Q. And is that for Los Angeles? 2.1 Α. Yes. 22 So at Los Angeles were you asked to Q. 23 perform the duties as deputy coroner with respect 24 to pathologic issues? 25 A. Yes, occasionally. 00058 Glancing quickly at the C.V. which was 1 2 mark as Exhibit 1, I noticed that you either edit 3 or review for perhaps about 27 peer review 4 journals; is that right? 5 Α. Yes. 6 Do many of those journals involve Q. 7 pathology? 8 Α. Yes. 9 Do they also involve cancer? Q. 10 Α. Yes. 11 And are you currently researching Q. 12 cancer? 13 Α. Yes. 14 And among your research interests, 15 Doctor, you mentioned before that you were 16 interested in this carcinoma bronchioloalveolar 17 lung carcinoma. Is that an interest that you've 18 had for some time? 19 A. Yes. 20 Are you currently researching on it? Q. 21 Α. Yes. 22 And you did describe for us, Doctor, Q. 23 the reason why you were interested in this is

24 because you noticed an increased incidence in BAC 25 over the years. 00059 1 When did you first begin seeing this 2 increased incidence? 3 Well, let me just back up a point. I 4 first became interested in BAC because it was a 5 tumor that was related to an extracellular matrix 6 that sometimes occurs in a scar or induced a scar 7 and was interested in extracellular matrix 8 interaction in breast cancer and BAC because it was 9 an unique cancer from that standpoint. And as I 10 was studying the matrix of BAC and the way BACs 11 metastasize I started to see an increase as I said before anecdotally in the number of cases and then 13 I asked the question is this really true or is it 14 just something I'm managing and that's when I 15 conducted this study. And since I was interested in disease 17 anyway, when I noticed that the disease was 18 increased in the incidence, I became even more 19 interested in trying to explain why this might be 20 so. And that's how I got interested in disease 21 even more. 22 Q. Doctor, I want to ask you some 23 questions about this rising incidence, but before I 24 do, let me get a little background here. Are all lung cancers the same? 25 00060 1 Α. No. 2 Are there different types or different Q. 3 kinds of lung cancers? 4 A. There are different types, there are 5 different cells of origin, and there's different 6 biologies. 7 And why is it important, Doctor, to Q. 8 know what type of lung cancer a patient may have? A. The biology may be different. The 9 10 prognosis may be different. The treatment may be 11 different. There are a number of reasons why it's 12 important to know. Is it also useful to know differences 13 Q. 14 in types of lung cancer as a researcher? Yes. A. 15 16 Q. And why is that? 17 Because since the types are different Α. 18 and the biologies are different, the molecular 19 events that cause or contributed to the cancer are 20 probably different and the etiologies are probably 21 different. 22 Now, when you say the etiologies of Q. 23 these different cancers are probably different, 24 what does etiology mean? 25 Α. Etiology means cause or causes. 00061 And you are currently researching a 1 2 potential etiology or cause of BAC; I do understand 3 that right? 4 Α. Yes. 5 And what is that potential cause or Q. 6 etiology of BAC? 7 It's the relationship of retroviruses Α.

8 or retroviral sequences to the etiology of BAC.

9 And that, Doctor, is ongoing research? Q. 10 Α. Yes. 11 Q. Being funded by various institutions? 12 It's being funded by the Margaret Α. 13 Early and this UC program. 14 Q. Doctor, what medical specialist or 15 which specialists is responsible for determining what type of lung cancer a patient may have? 16 17 Well, that would be a diagnostic Α. 18 pathologist. 19 And how do pathologists -- diagnostic 20 pathologists make these determinations about 21 specific lung cancer types? A. We take tissue that's been give to us 22 23 by a surgeon or pulmonologist, and we stain the 24 tissue and cut sections and look at the patterns 25 under a microscope. 00062 I want to again get to this rising Q. 2 incidence of BAC lung carcinoma. This is a study that you did back at 3 4 least prior to you publishing it, which it was 5 published in 1994? A. 6 Yes. 7 Q. And we've marked this now as Exhibit 8 9 Doctor, in that study, who was the 10 pathologist who reviewed the materials? 11 A. It was me. 12 Ο. And did you review all the pathology 13 that was listed in this case in this study? 14 A. Yes. 15 Q. Do you have the microscopic criteria 16 that you used for diagnosing BAC in this study? 17 Α. Yes. Is it listed in the paper that we've 18 Q. 19 marked as Exhibit 4, "Rising Incidence of 20 Bronchioloalveolar Lung Carcinoma"? 21 Α. Yes. 22 Q. Bronchioloalveolar lung carcinoma, 23 Doctor, is the same thing as BAC? 24 Yes. Α. 25 You have been using those terms Q. 00063 1 interchangeably? It makes your tongue tied to say 2. Α. 3 "bronchioloalveolar" many times. So it was 4 shortened to BAC. Some people use the term BAC. 5 Before we get too many terms here, 6 let's just stick with BAC. Now, doctor, is there 7 any clinical findings, and I know that you rely 8 upon the pathology here, but are there any clinical 9 findings in general that may be helpful in 10 identifying BACs? 11 Α. Well, BACs are associated with some 12 stereotypic type of clinical presentation. Of 13 course, I want to emphasize that the bottom line is 14 the tissue pattern under the microscope. So we can diagnose BAC even if the clinical presentation 15 16 isn't typical or we may not diagnose BAC. But the 17 typical presentation is a solitary peripheral 18 lesion of the lung that's pleural based that 19 puckers the pleura that may be associated with a

20 scar. That's the typical clinical presentation. Q. Doctor, are you familiar with the term 22 air bronchiograms? 23 Α. What are air bronchiograms? 24 Q. 25 That's a radiological term that refers Α. 00064 1 to a certain appearance on chest X-ray in which the 2 bronchus becomes prominent, and it's a finding that 3 radiologists make sometimes, and it's a finding 4 that is seen in BAC. Is it seen in BAC more than other 5 Q. 6 cancers? 7 Yes. And how does this phenomenon, this 8 Q. 9 finding, how does this happen? A. My opinion of the phenomenon is that 10 11 the BAC almost by definition grows a certain way in 12 the lung. It grows along alveolar septa and along 13 airways in a lepidic growth pattern and it fills 14 the alveolar lining. Alveolar, what is that? 15 Q. Those are the sacks that the lungs are 16 Α. 17 composed of that result in air exchange. When we 18 breathe, our lungs expand and then they retract, 19 and the airways are sort of like sponges. The 20 alveolar sponges are like the holes in a sponge. 21 Anyhow, the BAC fills these spaces 22 with proliferating cells that line the spaces and 23 they group in a lepidic or radial pattern. Q. They line the holes of these sponges? 24 25 Yes. They don't affect the bronchus. Α. 00065 And the bronchus is? 1 Q. The bronchus is a tube that conducts A. 3 air from our throat into the lungs and the bronchus serially divides like branches of a tree. BAC doesn't affect the bronchus, 5 6 unlike other cancers which can arise from the 7 bronchus. So when the BAC is involving the 8 alveolar spaces and the way we pointed out, it can 9 make the bronchus stand out on chest X-ray and 10 that's referred to as an air bronchiogram. 11 Q. Doctor, in the study that you did that 12 you were asked about before, that was published in 13 what journal? A. 1994. 14 15 And what journal, Doctor? Q. 16 "Cancer." Α. 17 Is that a peer review journal? Q. 18 Α. Yes. A. Yes.
Q. And is that the journal name of the 19 20 American Cancer Society? 21 A. Yes. 22 Q. You were the primary author in this 23 article? A. 24 Yes. 25 And is it the result of the research Q. 00066 1 that you did? 2 A. Yes. Now, Doctor, you've mentioned in the Q. 4 outset of the article -- and I'm going to give it

5 to you so you can have it -- bronchioloalveolar 6 lung carcinoma BAC is a form of lung cancer 7 exhibiting many features that distinguish it from 8 all other forms of lung cancer including non-BAC 9 adenocarcinoma. 10 You've already told us some of the 11 diagnostic things that make BAC different. Are there other things demographic or otherwise that 12 13 make BAC different from other lung cancers? 14 Α. Yes. For one thing, the biggest 15 demographic thing is that the female to male ratio 16 is close to 1 to 1. Most other lung cancers still 17 have a very high male to female ratio. Could you explain what that means? 18 Q. 19 Α. Well, it's just the number of cases of 20 a particular lung cancer in males versus females. 21 So there's a lot more males coming 22 down with these squamous cell carcinoma or 23 adenocarcinomas than females generally? 24 Α. Yes. But in BAC, what's the relationship? 25 Q. 00067 A. Like I said, it's about a 1 to 1 1 2 female to male ratio. So, in other words, we see 3 about as many cases of BAC in females as we do in 4 males, and that ratio has been relatively constant 5 over the years. Even while BAC has been rising 6 Q. 7 dramatically? 8 Α. Yeah, it's been rising in both sexes 9 about equally. 10 Q. And is there, Doctor, anything else 11 that makes BAC different or unique? 12 A. Well, I said before, it tends to be 13 peripherally. It tends to arise in relationship to 14 a scar, which it either can antedate or induce. 15 We do see it in a high percentage of 16 nonsmokers. It also tends sometimes to occur in a 17 multifocal manner. 18 Doctor, could you describe the cases 19 that you reviewed for this study? How did you get 20 those cases? Well, I just did a computer retrieval 21 22 of every case of lung cancer that existed at our 23 institution since the mid-'50s. 24 And, Doctor, all of these cases had a Q. 25 diagnosis; is that right? 00068 1 Α. Yes. 2 So why was it that you rediagnosed 3 them? 4 Well, because sometimes diagnostic Α. 5 criteria change over the years or different people 6 are involved and the pathologist who reviewed the 7 cases in the 1950s, most of those would not be 8 around, and I wasn't there in the 1950s. So if you 9 have different people reviewing cases, that 10 introduces what's called an intraobserver bias, if 11 you will. 12 So that's why if you do any study of a 13 retrospective nature you want to make sure that you 14 or your associates or both review all the cases so 15 you can have an uniform criteria that you apply.

So did you apply a uniform criteria to 16 Ο. 17 all 1,527 cases of lung cancer? 18 A. Yes. 19 And, Doctor, when you did this Q. 20 rereview and rediagnosis with this uniform criteria 21 that you employed, how much variability did you 22 find from the original diagnoses? 23 A. Surprisingly, there wasn't that much 24 variability. Around 5 percent or so. 25 Q. Doctor, the criteria for diagnosing 00069 1 BAC, is that contained in the part of the article 2 called "Diagnostic Criteria"? 3 Α. Yes. 4 Q. And rather than me read those couple 5 paragraphs and ask you to explain it, let me ask 6 you a couple of questions. 7 Do you always use the criteria that 8 you have here in diagnosing BACs? 9 Α. Yes. Do you use them in your practice as a 10 Q. 11 pulmonary pathologist at UCLA weekly? A. Yes. 12 13 Roughly on average how many lung Q. 14 cancers do you see weekly? 15 A. I would say between 5 and 20. It can 16 range. It could be a great range. 17 Q. And did you apply the same criteria 18 that you've listed here in this study to the case 19 that we brought you to look at, Mrs. Boerner? A. 20 Yes. 21 Now, Doctor, earlier you were asked Q. 22 some questions about some of the photomicroscopy 23 that you took in this case, and did you, in fact, 24 take pictures of the pathology of the Mrs. 25 Boerner's case? 00070 1 Α. And let me just show you what has been 2. 3 marked previously as 3-A and, again, Doctor, could 4 you tell us now that you have it in front of you 5 there, could you tell us what it is that's 6 significant in that picture and what it is about 7 that picture that leads you to concludes Mrs. 8 Boerner's cancer was a BAC? 9 A. First of all, I see neoplastic cells. 10 I see malignant epithelial cells that are lining 11 the alveolar spaces and growing along them. They 12 are growing along them in a single cell fashion. 13 They are not undergoing proliferation. 14 They are preserving the interstitium 15 framework of the lung. That's intact. They're not 16 destroying it. They are not invading into it. 17 They are just invading along this interstitial 18 framework. This is what is called a lepidic growth 19 pattern and that's what I used to diagnose BAC. 20 Doctor, you refer in your article to 21 what is called classic BAC. How would you 22 characterize what Mrs. Boerner's cancer exhibits? 23 Well, in the areas of BAC, they were 24 classic. They were nonmucinous, as I said. The 25 cells looked a little bit like Clara cells, 00071

```
1 although without immunio or EM, I couldn't rule out
 2 a type 2 pneumocytes.
 3
          Q. And what's the significance of those?
 4
               Those are two cells that one finds in
 5 the terminal bronchi and the alveoli that are
 6 thought to give rise to BAC. So it's those cells
   that are transferred that produce BAC.
7
 8
          Q. Now, Doctor, in your article you make
9
   the statement, and this is again the article that
10 was referred to earlier by Plaintiffs' counsel
11 Exhibit No. 4. You make the statement:
                "Furthermore, the degree of
12
13
          nuclear pleomorphism or the degree of
14
          intraalveolar proliferation, including
15
          papillary proliferation manifested by
16
          the carcinoma cells, did not prohibit
          the assignment of the tumor into the
17
          BAC category if there was significant
18
19
          growth in a single cell pattern along
20
          alveolar septa."
                What does that all mean, Doctor, in
21
22 plain English?
23
          Α.
               What that means, if you have a
24 significant degree of cell division or
25 proliferation, that can produce a pattern where you
00072
1 no longer have single cells growing in a lepidic
 2 growth pattern. And if those cells make a granular
 3 pattern you might make the diagnoses of
 4 adenocarcinoma of a non-BAC type.
 5
                What I'm stating here is, irrespective
 6 of whether there was proliferation in some areas of
 7 some of these cases, if I saw a classic BAC pattern
8 in a reasonable area of the tumor, I would conclude
9 that that cancer belonged in the BAC category.
          Q. And you applied that to all 1,527
10
11 cases you reviewed?
          A.
12
               Yes.
13
                Likewise, doctor, it goes on to say in
          Ο.
14 your article:
15
                 "The presence of solid areas of
          moderately to poorly differentiated
16
17
          adenocarcinoma also did not exclude the
18
          tumor from the BAC category if areas of
19
          classic BAC were present."
20
          Α.
               Yes.
21
          Q.
                Now is that the same type of thing
22 that you just described?
23
          Α.
                Yes.
24
                So even if you saw areas that were of
25 another poorly differentiated cancer type, if you
00073
1 saw classic areas of BAC, how would you diagnose
 2 it?
 3
                I would conclude it was a BAC that
 4 underwent dedifferentiation into a more aggressive
 5 sub type.
 6
                Doctor, you talk about
          Q.
 7
   dedifferentiation. In fact, let me just read to
 8 you from the next column on page 1164 of your
 9 article on cancer.
10
                "Cases accepted as BAC were
11
          evaluated for the presence of areas of
```

dedifferentiation into more solid 12 13 moderately to poorly differentiated adenocarcinoma." 14 15 What is "dedifferentiation," Doctor? 16 Dedifferentiation is a form of tumor 17 progression that many cancers exhibit. Not just 18 lung cancers. As cancers evolve, they are thought 19 to acquire more molecular abnormalities, more 20 instability of their gene form that contributes to 21 what we determine dedifferentiation. It's a 22 conversion to a more aggressive sub type, and it's 23 a general pathway of progression that many cancers 24 exhibit as they progress, as they metastasize, 25 et cetera. 00074 1 So dedifferentiation, Doctor, means Q. 2 that the cancer becomes less defined; is that 3 right? I would say less differentiated. In 5 other words, the cancer becomes very aggressive, 6 very deranged. It less resembles its normal counterpart. It's less differentiated. A normal 7 cell is considered a differentiated cell. 8 Q. Is classic BAC a well-differentiated 9 10 pattern? 11 Yes. It's considered well 12 differentiated. Q. Now, can it dedifferentiate from a 13 14 poorly differentiated pattern to a well 15 differentiated pattern? That's not what's seen in the vast 16 Α. 17 majority of cancers. The pathway is one 18 direction. It begins as well differentiated, it 19 progresses to poorly differentiated rather than the 20 reverse. 21 When you reviewed the cases for your 22 study that we've marked as Exhibit 4, those 1,527 23 cases, what percentage roughly of BACs did you see 24 that had patterns of dedifferentiation? 25 Α. I think overall it was around 20 00075 1 percent, but the percentage varied depending on the 2 specific kind of BAC. In the nonmucinous type, it 3 was around 10 percent. 4 Q. You also wrote in your article, 5 Doctor, about BACs and association with scarring. 6 Do you recall that? 7 A. Yes. 8 And you've also written an article Q. 9 about lung cancers that can cause desmoplasia. Do 10 you remember that article? 11 Yes. Α. 12 So you've published on this issue in Q. 13 the past? 14 Α. Yes. 15 What is "desmoplasia," Doctor? Q. 16 Desmoplasia is the formation of 17 fibrous tissue, scarring if you will, as a result 18 of tumor invasion. So there are cancers such as 19 breast cancer which induce a profound scar and 20 produce is a hard lump in the breast. BAC is 21 another cancer that can induce a scar, which we 22 term desmoplasia.

```
Q.
               You've used a few terms. Fibrosis is
23
24 fibrosis. And scarring, for all practical
25 purposes, the same thing in the lung?
00076
1
          Α.
               Yes.
2.
              And fibrosis or scarring, can that be
          Q.
3 caused by cancer?
          A.
4
5
          Q.
               Can other things cause fibrosis and
 6 scarring?
7
         A. Yes.
              When the fibrosis and scarring is
8
          Ο.
9 caused by cancer, what's that called?
         A. Desmoplasia.
10
11
          Q.
               In the article that you wrote that
12 maybe we should cite it for the record here was
13 "The Extracellular Matrix of Pulmonary Scar
14 Carcinoma is Suggestive of a Desmoplastic Origin."
15
               Do you recall that article?
16
               Yes.
          Α.
          Q. You've been studying desmoplasia since
17
18 at least 1986? And perhaps some years before?
19
          A. Yes.
20
          Q.
               In that area, do you discuss the
21 difference between scars that are caused by cancer,
22 those desmoplasia scars and scars that are not
23 caused by cancer?
24
         A.
               Yes.
               And the scars that are not caused by
25
00077
1 cancers, do you refer to those as noncarcinoma
2 scars?
3
               Yes.
          Α.
               What are some of the causes of
4
         Q.
5 cancers -- strike that.
              What are some of the causes of
 6
7
   noncarcinomative scars?
          A. Well, in the lung, common causes are
8
9 infarcts, old tuberculosis or histoplasmosis.
10 That's called granulomatis disease. Sometimes
11 trauma to the lung could be a cause of an old
12 scar. Sometimes there's just a scar in the pleura
13 that we don't know the reason why it occurred but
14 we do find them.
15
          Q. And, Doctor, can you tell when you
16 look at a scar that's a granuloma, can you tell
17 that scar is a granuloma scar and not a scar from a
18 cancer when you look at it under a microscope?
19
               Most of the times we can.
          Α.
20
               And when you see an infarct, Doctor --
          Q.
21
                First of all, what is an infarct?
22
               An infarct is death, necrosis of
          Α.
23 tissue due to poor blood supply, and it's usually
24 due toe an occlusion in a blood vessel, which is
25 called the thrombus or an embolus.
00078
               Do you see these infarcts in a wide
1
          Q.
 2 range of patients?
          Α.
 3
               Yes.
 4
          Q. Do you see them in smokers?
 5
          A. Sometimes.
 6
          Q. Do you see them in nonsmokers?
 7
                Yes.
          Α.
```

Is there anybody who has ever said Q. 9 that infarcts are in any way related to smoking? A. Not pulmonary infarcts. 10 11 And that's what we're talking about in Q. 12 this case is pulmonary infarcts. Infarcts that are 13 pulmonary infarcts are infarcts in the lung? A. They are quite common. We see them 14 15 all the time, incidentally, at autopsy for example. 16 Q. Doctor, how can you tell if the scar 17 is an infarct when you look at it under a 18 microscope? In the lung I said that an infarct was 19 20 defined as necrosis or death of tissue. One type 21 of tissue that we have in our lungs is called 22 elastic tissue, and it's the reason why our lungs 23 are so resilient. They expand and contract when we 24 breathe. The elastic tissue which is in a sense 25 very similar to a rubberband, for example, in terms 00079 1 the its recoiling properties, is very resistant to 2 necrosis and low blood flow. So it remains, it's 3 preserved. Because the other tissue is destroyed 4 in an infarct, the elastic tissue collapses on 5 itself. So if you see a scar that has a lot of 6 elastic tissue jumbled together, you would conclude 7 that was an infarct. 8 Doctor, this elastic tissue, does it Ο. 9 have a certain appearance that you can describe 10 under a microscope? 11 Α. Yes. 12 What is that appearance? Q. 13 A. It looks very serpentinus. It's 14 serpentine like, with coils. Is it wavy? 15 Q. 16 Α. Yes, it's wavy. When you say serpentine, is that wavy 17 Q. 18 and jumbled up? 19 Α. If you've ever been to a zoo and seen 20 snakes, you can see where the term "serpentine" 21 comes from. In addition to the research that Q. 23 you've done, including the article on the 24 extracellular matrix of pulmonary scar carcinoma, 25 in addition to that research, are you familiar with 08000 1 pathology literature that discusses whether scars 2 cause cancer or cancer causes scars? A. Yes. 3 4 Q. And are you familiar that there have 5 been various opinions raised about this issue over 6 the past 20 years or so? A. Yes.Q. Doctor, in your opinion with a 7 8 9 reasonable decree of medical certainty, do you 10 believe that scars cause cancer or that cancer 11 causes scars or that both occur in the lung? A. I think under certain situations both 12 13 can occur or at least a scar can predispose to a 14 cancer or the cancer can induce the scar. 15 Q. And is that also true for BACs? Can 16 BACs arise from scars and can scarring -- strike 17 that. 18 Can BACs arise from scars or can BACs

19 cause scarring? 20 A. Both can occur. 21 Q. Are you familiar with the epidemiology 22 literature that has demonstrated certain lung 23 cancers are associated with smoking? 2.4 Yes. Α. 25 Q. And when researchers say that there is 00081 1 an association between a certain type of lung 2 cancer and smoking, does that mean that there is a 3 higher incidence in smokers when compared to never 4 smokers? 5 Α. Yes. If smoking caused a certain type of 6 7 lung cancer, would you expected to see an 8 association between smoking and that lung cancer? 9 A. I would expect to see more than just 10 an association. It would have to be a strong 11 association, and it would also have to fulfill 12 certain epidemiological criteria such as a 13 relationship between dose of exposure and incidence of disease. There's a number of criteria that have 14 15 to be met before an association becomes causative. 16 Q. Even with epidemiological principles? 17 Α. Especially with epidemiological 18 study. 19 Now, so therefore, one of the things Ο. 20 among others you would expect to see is a higher 21 incidence or more of a particular type of lung 22 cancer in smokers than in nonsmokers before you 23 could say epidemiologically that the smoking caused 24 the type of lung cancer? 25 They would have to at least be that. Α. 00082 In your research, Doctor, and in the 1 2 study you published in 1993 or 1994, the "Rising Incidence of Bronchioloalveolar Lung Carcinoma and 4 it's Unique Clinicopathological Features, " in that 5 study, Doctor, was there an association between BAC 6 and smoking? 7 Our cases of BAC were sort of roughly 8 equally divided into three groups. About a third 9 never smoked, a third infrequently or remotely 10 smoked, and a third smoked. So based on that 11 distribution, there was not a convincing 12 association of smoking with BAC. 13 In fact, Doctor, was there any 14 increase in BAC in smokers compared to the 15 nonsmokers? 16 A. No BAC increased in both smokers as 17 well as nonsmokers. 18 Q. And, Doctor, this is based upon a 19 study using the clinical criteria for BAC that you 20 used in this case? 21 Α. Well, the pathological criteria that I 22 used. 23 Let me get the question correctly. 24 This finding that there was no 25 increase in BAC in smokers compared to nonsmokers 00083 1 utilized and was based upon the pathological 2 criteria for bronchioloalveolar lung carcinoma that 3 you applied in this case to Mrs. Boerner?

4 Yes. Now, in doing the research you have 5 Q. 6 done on BAC, Doctor, have you reviewed the 7 published scientific literature? 8 A. Yes. 9 And are you aware of any scientific Q. evidence that shows there is an increased incidence 10 of BAC in former smokers when compared to smokers? 11 12 A. There are studies that show an 13 association in former smokers with adenocarcinoma 14 but not specifically. But the association -- the 15 studies that addressed a relationship of smoking to 16 BAC, while there are sometimes some associations, they are not strong. They are not convincingly of 17 18 a strong enough nature to imply a causation. 19 Q. Referring to the lung cancers that 20 arise from a scar, do you know of any research that 21 has associated those cancers with smoking? 23 Q. I want to talk a little wee bit about 24 Mrs. Boerner's pathology and you've already told us 25 that you've reviewed the pathology, and you've told 00084 1 us you've reviewed the slides from the biopsy 2 lobectomy from the metastasis and the neostatial 3 nodes. And I understand from the review of all 4 this pathology that your diagnosis with a 5 reasonable degree of medical certainty was a 6 bronchioloalveolar lung carcinoma with 7 dedifferentiation; is that correct? 8 A. Yes. 9 Q. And, again, the diagnosis in Mrs. 10 Boerner's case and coming up with that diagnosis, 11 you applied the same criteria that is listed and 12 set forth in your article that we've attached as 13 Exhibit 4; correct? 14 Yes, and also the same criteria I Α. 15 would apply to any case that I saw in my duties as 16 a pathologist at UCLA. 17 If the pathology from Mrs. Boerner's 18 cancer had been in the population of cancers that 19 you reviewed for your study that was reported in 20 the journal "Cancer," what lung cancer type would 21 she been classified as? 22 A. I classified her as a nonmucinous BAC 23 with areas of dedifferentiation. Now let's talk about the 24 25 dedifferentiation. What did the areas of 00085 1 dedifferentiation resemble? 2 There were areas that looked like 3 adenocarcinoma, squamous carcinoma. There were 4 areas that were kind of mixed adenosquamous and 5 there were areas of clear cell carcinoma. 6 Q. And is that reflected in what has been 7 marked as Exhibit 3 B? 8 A. Yes. 9 And, doctor, 3-B is a picture that was Q. 10 taken from a slide of Mrs. Boerner's pathology? 11 Yes. Α. To an reasonable degree of medical 12 13 certainty, Doctor, did the dedifferentiated portion 14 of Mrs. Boerner's tumor develop after the BAC?

```
15 A. Yes.
```

00087

16 Q. To a reasonable degree of medical 17 certainty, Doctor, did the dedifferentiated portion 18 of Mrs. Boerner's tumor develop after 1981?

19 A. Yes. I feel that the dedifferentiated 20 tumor is a very rapidly growing, very aggressive 21 tumor that could not have been around very long, 22 and so I wouldn't think that it could have existed 23 for, you know, a period of 15 years. Therefore, it 24 would have had to develop after 1981.

Q. And did smoking have anything to do 00086

with the dedifferentiation of Mrs. Boerner's tumor?
Given the fact that she was no longer

3 smoking after 1981 and given the fact that this 4 tumor is a BAC, I'm not convinced that there's a 5 link between the two.

Q. Did you see BAC tumors, Doctor, in your study that you have published on the rising incidence of bronchioloalveolar lung carcinoma and it's unique clinicopathologic features that had dedifferentiated into areas of squamous and adeno and adenosquamous and clear cell like you've seen some in Mrs. Boerner's case?

13 Α. The majority of cases of 14 dedifferentiation were those into adeno. Poorly 15 differentiated adeno or undifferentiated. The 16 purpose of that study, the main thrust was to discuss and to note the rising incidence of BAC. 17 18 It wasn't a study intended to analyze this very 19 specific type of dedifferentiation, per se, and so 20 I don't -- the study doesn't mention squamous 21 differentiation or clear cell. I'm sure if I went 22 back looked specifically for that. I would 23 probably see some case that's would fit that. But 24 the ones that were mainly seen were those into 25 poorly differentiated adeno.

1 Q. Is there any or evidence besides the 2 microscopic appearance that support your view that 3 this is a BAC with dedifferentiation?

That's a major finding, but the fact 4 5 that there's a scar, that the tumor is arising 6 peripherally, that there are precursor lesions of 7 bronchioloalveolar metaplasia and hyperplasia and 8 hyperplasia with atypia that seem to be progressing 9 into the areas of BAC, the lack of any change in 10 the major bronchi of a displastic or metaplastic 11 nature, and the fact that there's a big scar with elastic tissue right next to where the BAC is, all 12 13 those things are supportive of that diagnosis, but 14 the major finding is the lepidic growth pattern and 15 the finding of BAC-type cells.

16 Q. Let's talk a minute about these 17 precursor lesions. Doctor, what are precursor 18 lesions?

19 A. In humans, cancer is not thought to
20 begin and develop overnight. It's not a single hit
21 where a normal cell becomes a frankly malignant
22 cell. Human cancers are thought to go through a
23 series of steps sort of like going up an
24 escalator. You start at the bottom; your cell is
25 normal. You get to the top and you have a cancer,

00088 1 but on the way, you have to go through a series of 2 changes or steps, and those steps are defined as 3 "precursor lesions." They are changes that have 4 occurred in cells that are not cancer yet, but they 5 are developing along those lines. And precursor 6 lesions in the lung, there's really two kinds of 7 precursor lesions. There are those that arise in 8 the central airways that are called bronchial 9 metaplasia and carcinoma and cito and those that 10 arise peripheral that are called bronchioloalveolar 11 metaplasia and metaplasia. Q. You said the ones that arise in the 12 13 bronchi are the precursors. Are those like the air 14 tubes of the lungs? 15 Α. Yes. 16 And you say those give rise to cancer Ο. 17 eventually? 18 Yes. 19 Ο. Are those precursor lesions associated 20 with smoking? A. Oftentimes they are.Q. And have you done study on precursor 21 Q. 22 23 lesions? Are you currently doing studies? 24 A. Yes. 25 Did you see any of those Q. 00089 1 smoking-related precursor lesions in Mrs. Boerner's 2 cancer? 3 Α. No. 4 In fact, Doctor, do you have some Q. 5 pictures of where those precursor lesions would be 6 to show us and to show us what the effect on 7 Mrs. Boerner's cancer would be? A. I have two pictures that are labeled C 8 9 and D of the normal appearing bronchi. In fact, 10 one picture is adjacent her cancer but doesn't show 11 any changes. 12 Now, let's talk about what we have as Q. 13 4-C first. Let me look at 4-C. 4-C is a section of a major bronchus. 14 15 Now, point out to us if you would, and 16 you are going to have to maybe draw with a pen on 17 this an arrow to where the bronchus really is. 18 What is the bronchus? 19 It's this structure which is lined by Α. 20 these cells. Q. Could you just put an arrow on that 22 picture? 23 Sure. (The witness complies.) Α. 24 Now, if there were precursor lesions Q. 25 due to smoking, this is where you'd expect to see 00090 1 them in this bronchial-type tissue? 2 Α. Yes. 3 And do you see any precursor lesions Ο. 4 there at all? 5 No, in fact, the bronchi look totally 6 healthy, totally normal. They have cilia. 7 Q. Is this representative of the bronchi 8 that you saw in Mrs. Boerner's pathology? 9 Α. 10 Doctor, what do we have as 4-D? Q.

```
It's another section of a bronchus.
11
          Α.
12 The only difference is there's some tumor near by.
13
         Q. Now, what I'd like to you do, again,
14 if you would, I hate to draw on your pictures but
15 just put an arrow to the tumor, if you would.
16
               (Indicating.)
17
               And put a little "T" there if you
18 would.
          A. (Indicating.)Q. Now that's tumor. Is that tumor at
19
20
21 the bronchus?
          A. It's under the bronchus.
22
              Did you see any tumor, Doctor, that
2.3
          Q.
24 was arising from the bronchus?
25
          Α.
               No.
00091
1
               Is this tumor arising from the
          Q.
2 bronchus?
3
         Α.
               No.
 4
          Q.
              Now, where is the bronchus?
               It's overlying the tumor.
 5
          Α.
 6
               Could you again put an arrow to that.
          Q.
               (Indicating.)
 7
          Α.
8
               And what do we see with that bronchus,
          Q.
9 Doctor?
10
               The bronchus looks normal. There's
          Α.
11 cilia, and it's normal in appearance.
               Again, is this the area where you
12
13 would expect to find the smoking-related precursor
14 lesions if they were to develop?
15
               Yes.
          Α.
16
               And you saw no precursor lesions
          Q.
17 related to smoking?
18
               Right. And I also saw no precursor
19 lesions going into the tumor from the vantage point
20 of the bronchus.
          Q. And, Doctor, how would you define this
21
22 bronchus and that cilia?
         A. Normal.
23
24
          Ο.
               What is cilia?
25
              Cilia is a structure that's related to
          Α.
00092
1 what's seen in certain cells, especially
 2 respiratory cells, and it has a flagellar-like
 3 function. These are structures on the outside of
 4 the cell that are thought to move difficult
 5 particles that go into our lung. It's thought to
 6 sort of move them out of our lung.
 7
               Like a little sweeper?
          Q.
 8
          Α.
               Yes.
              And these little cilia, they are like
9
          Q.
10 little tiny, tiny, tiny hairs?
11
          A. Yes.
12
          Q.
               And they are very, very delicate?
13
          A.
               Yes.
14
               And they are in absolutely normal
          Q.
15 shape in this picture?
16
          Α.
               Yes.
17
          Ο.
               And is that what you saw throughout
18 the pathology of Mrs. Boerner?
19
          A. Yes, in her bronchus.
               Again, the bronchus, again, are the
          Q.
21 air tubes where the cancer did not arise?
```

22 Α. Right. 23 Now, Doctor, did you see any precursor Q. 24 lesions in Mrs. Boerner's case? 25 A. Yes. They were in the peripherally --00093 1 they were peripheral lesions, and I had termed them 2 bronchioloalveolar metaplasia, hyperplasia and hyperplasia with atypia. 4 And what is, again, the significance Ο. 5 of finding these precursor lesions in 6 Mrs. Boerner's cancer? These precursor lesions are thought to 7 8 antedate the development of BAC, and finding them 9 was significant because it told me that that's 10 where the BAC was taking origin from. In addition, 11 these precursor lesions are not well understood and 12 their genesis is unknown and, again, these are 13 precursor lesions who's progression has not been 14 linked to smoking. 15 Q. You said it was important to know 16 where these precursor lesions were. Where were 17 they? They were adjacent to the scar in the 18 19 periphery of the lung and adjacent the developing 20 BAC. 21 So if I can get a picture of this, 22 Doctor. We have this big scar that you said was 23 from an infarct, and then from that scar we have 24 these precursor lesions, and then the precursor 2.5 lesions developed into the BAC cancer of 00094 1 Mrs. Boerner? A. Yes. 3 Q. Do you have any pictures where you can 4 take us through this process and show us how this 5 looked? 6 Yes, they are labeled Boerner G, H and Α. 7 I and Y. If you would, Doctor, I'm just going 8 Q. 9 to let you take us through those one at a time, and 10 if you could kind of start with tell us what we 11 have and what it looks like. A. Well, we'll start with G. G shows 12 13 some bronchioloalveolar metaplasia in the alveolar 14 spaces. 15 Alveolar spaces, again, are what? Q. They are the sacs or the sponge areas 16 Α. 17 of the lung that fill with air. 18 And this bronchioloalveolar metaplasia Q. 19 did you say? 20 Α. Yes. 21 Is that cancer? Q. 22 Α. No. 23 Q. What is it? 24 A. It's a switch from the alveolar cell 25 which normally lines the alveolar space to a 00095 1 bronchioloalveolar cell. It's a cell that 2 resembles a cell that's normally in the bronchus. 3 Q. This is a cell that's undergone a 4 change but not cancerous? 5 A. A metaplastic change. 6 And metaplastic means? Q.

Metaplastic is a switch from a mature Α. 8 adult cell type to a mature adult cell type. It 9 may be or may not be a precursor to cancer, but 10 it's not cancer. 11 Q. But it gives you as a pathologist some 12 concerns about what's going on here? A. Especially if you see the next lesion 13 14 which is bronchioloalveolar hyperplasia. Q. And what is that? 15 It's the same beginnings as what I 16 A. 17 described for metaplasia. It's a switch from the 18 normal alveolar cell to the bronchiole cell, but 19 the bronchiole cell has undergone some 20 proliferation cell division. It's more numerous 21 and thicker in terms of its appearance. 22 Q. Now what slide is that what we're 23 looking at? A. That's Boerner H -- excuse me. H. 24 25 So we've moved now from the bronchiole Ο. 00096 1 metaplasia. We're getting more transformation here 2 but we're not cancer yet? 3 A. Correct. 4 Q. And as this continues to move, Doctor, 5 what's the next event? 6 A. The next molecular event is unknown 7 but the next morphological description --Q. What is morphological description? 8 It's what pathologist see under the 9 Α. 10 microscope. The next thing we see is in Boerner I, 11 and again, the process has continued in its 12 degree. There's more proliferation. There's more 13 involvement of the alveolar space. There are more 14 bronchial cells. They are more prominent, more 15 numerous and more atypical in terms of their size 16 and nuclear features. 17 Q. And, again, these are all taken from 18 Mrs. Boerner's pathology? 19 A. Right. 20 What's the next picture you have Q. 21 there? The next picture is labeled Boerner J, 22 23 and it's an area where BAC has now developed from 24 one of these foci. 25 Q. And these foci being the what? 00097 1 These foci of bronchioloalveolar Α. 2 hyperplasia and atypia. 3 So now we have seen the transition 4 from the bronchio metaplasia all the way through to 5 finally it reaches frank cancer? A. 6 Yes. 7 We have BAC cancer? Q. 8 Α. Yes. 9 Q. Doctor, if you could just go back to I 10 for a minute and what do we have with I? I is a focus of bronchioloalveolar 11 Α. 12 hyperplasia with atypia. 13 Q. And what is atypia? 14 Again, it's a descriptive 15 morphological finding that connotes variation in 16 shape and size of the cells and the nuclear 17 features.

Does atypia mean it's more abnormal 18 Q. 19 than the material we saw before? 20 A. Yes. 21 So this is, again, becoming more and Q. 22 more closer to becoming cancerous? 23 Yes. Doctor, the remaining two slides, what 24 Ο. 25 do you have there? 00098 1 These are slides that are labeled E Α. 2 and F. Again, from Mrs. Boerner's pathology. 3 E shows this pulmonary scar, which is 4 5 filled with a very pink wavy serpentinus elastic 6 7 Serpentinus, is that the same thing as Q. 8 that serpentine, the snake-like wavy things you 9 talked about? 10 Yes. 11 And could you tell us, is this high 12 magnification, low magnification? 13 A. Sort of middle. How big was this scar? Q. 14 It was fairly obvious. I would say it 15 Α. 16 measured at least a few millimeters. 17 Q. Is it something that one could easily 18 miss on reviewing the pathology? 19 A. I don't think so. It was very 20 prominent when I reviewed these slides. In fact, I 21 also note that when I reviewed the records that 22 when this tumor was removed, the person who 23 described it grossly -- you know, I didn't have an 24 opportunity to do since I just saw the slides. The 25 person who reviewed it describes an area of pleural 00099 1 thickening and describes the fact that the tumor was arising subpleurally. I bet you any money that 3 that area of pleural thickening is what corresponds 4 to this scar that I'm seeing. 5 Q. This infarct scar? 6 Α. Yes. 7 And, Doctor, is there any doubt in Q. 8 your mind whether that scar was caused by the 9 cancer? 10 A. Oh, the scar preceded the cancer. 11 scar was due to an infarct. 12 The last picture that you asked me 13 about, which was F, in fact shows the scar on one 14 side and the BAC on the other. So it shows the 15 juxtaposition of the two processes. 16 And is that a low power? Q. 17 Α. Yes, it's a low power. Doctor, the slides that you have taken 18 Q. 19 that shows the transition that we have gone through 20 from the bronchioloalveolar metaplasia to the 21 atypia and into the frank cancer, where were these 22 in relationship to J? 23 A. Well, they are in the vicinity. 24 mean, J doesn't show the foci metaplasia because 25 you can only show so many things on a given 00100 1 picture. The point of J is to show the scar and 2 the BAC. But the foci are around the scar in

```
3 different areas.
 4
        Q. In your opinion, Doctor, with a
 5 reasonable degree of medical certainty, did
 6 Mrs. Boerner's lung cancer arise out of a
7 preexisting infarct scar?
8
          Α.
               Yes.
               In light of your findings that Mrs.
9
          Q.
10 Boerner's cancer was a bronchioloalveolar lung
11 carcinoma and was arising from a preexisting scar,
12 can it be said with a reasonable degree of medical
13 probability that smoking was a cause of
14 Mrs. Boerner's lung cancer?
15
          Α.
               No.
16
               Can you say with a reasonable degree
          Q.
17 of medical probability that if Mrs. Boerner had
18 never smoked she would have avoided her lung
19 cancer?
20
         Α.
               No.
21
         MR. SHEFFLER: Okay. I'm through. Hello?
22
         MS. HARTLEY: I don't have anything else.
23 Is he going to read?
          MR. SHEFFLER: Yes, he is.
24
25
          MS. HARTLEY: Okay. We'll see you later
00101
1 then. Have our court reporter get us a copy.
    MR. SHEFFLER: One thing, we may try to --
3 what we have for the purposes of the deposition are
4 some the pictures printed off. Now these are not
 5 quite as legible as they would like because of the
 6 printer that was used and we may reprint them.
 7 we do, Dr. Barksy will make the arrows, et cetera,
8 as he did today and we will supply you with that
9 copy.
          MS. HARTLEY: Okay. Thank you.
10
          (Ending time: 12:11 P.M.)
11
12
13
14
15
16
17
18
19
20
21
22
23
24
25
```